

- 2-cyclopentyl-1,2-dihydro-4-quinazolinamine;
1,2-dihydro-2,2-dimethyl-4-quinazolinamine;
2-ethyl-1,2-dihydro-2-methyl-4-quinazolinamine;
1,2-dihydro-2-methyl-2-phenyl-4-quinazolinamine;
5 2-(2-furyl)-1,2-dihydro-4-quinazolinamine;
1,2-dihydro-2-(2-thienyl)-4-quinazolinamine;
1,2-dihydro-2-(4-pyridyl)-4-quinazolinamine;
1,2-dihydro-2-(1H-imidazol-2-yl)-4-quinazolinamine;
1,2-dihydro-2-(2-thiazolyl)-4-quinazolinamine;
10 2-(4-cyanophenyl)-1,2-dihydro-4-quinazolinamine;
2-(4-dimethylaminophenyl)-1,2-dihydro-4-quinazolinamine;
1,2-dihydro-2-(4-nitrophenyl)-4-quinazolinamine;
2-(9-anthracenyl)-1,2-dihydro-4-quinazolinamine;
2-(4-amino-1,2-dihydroquinazolin-2-yl)benzenemethanol;
15 1,2-dihydro-2-(2-nitrophenyl)-4-quinazolinamine;
1,2-dihydro-2-(5-nitro-2-thienyl)-4-quinazolinamine;
ethyl 2-(4-amino-1,2-dihydroquinazolin-2-yl)-1H-pyrrole-1-carboxylate;
1,2-dihydro-2-(trimethylsilylethynyl)-4-quinazolinamine;
spiro[cyclopentane-1,2'(1' H)-quinazoline]-4'-amine;
20 spiro[cyclohexane-1,2'(1' H)-quinazoline]-4'-amine;
5-chloro-2-(2-furyl)-1,2-dihydro-4-quinazolinamine;
5-chloro-1,2-dihydro-2-(2-thienyl)-4-quinazolinamine;
5-fluoro-1,2-dihydro-2-phenyl-4-quinazolinamine;
5-fluoro-2-(2-furyl)-1,2-dihydro-4-quinazolinamine;
25 5-fluoro-1,2-dihydro-2-(2-hydroxyphenyl)-4-quinazolinamine;
5-fluoro-1,2-dihydro-2-(3-hydroxyphenyl)-4-quinazolinamine;
5-fluoro-1,2-dihydro-2-(4-hydroxyphenyl)-4-quinazolinamine;
ethyl 3-(4-Amino-5-fluoro-1,2-dihydroquinazolin-2-yl)-1H-pyrrole-1-carboxylate;
5-fluoro-1,2-dihydro-2-(2-thienyl)-4-quinazolinamine;
30 5-fluoro-1,2-dihydro-2-(2-thiazolyl)-4-quinazolinamine;

- 5-fluoro-2-(4-fluorophenyl)-1,2-dihydro-4-quinazolinamine;
5-fluoro-1,2-dihydro-2-(4-methoxyphenyl)-4-quinazolinamine;
5-fluoro-1,2-dihydro-2-(4-(methylthio)phenyl)-4-quinazolinamine;
5-fluoro-1,2-dihydro-2-(2-(trifluoromethyl)phenyl)-4-quinazolinamine;
5 5-fluoro-1,2-dihydro-2-(4-(trifluoromethyl)phenyl)-4-quinazolinamine;
5-fluoro-1,2-dihydro-2-(1-methylethyl)-4-quinazolinamine;
2-cyclobutyl-5-fluoro-1,2-dihydro-4-quinazolinamine;
5-fluoro-2-(2-furyl)-1,2-dihydro-2-methyl-4-quinazolinamine;
2-(2-furyl)-5-(methylthio)-1,2-dihydro-4-quinazolinamine;
10 1,2-dihydro-1-methyl-2-phenyl-4-quinazolinamine;
2-cyclopropyl-1,2-dihydro-1-methyl-4-quinazolinamine;
4-amino-1,2-dihydro-2-quinazolinepropanamine;
4-amino-1,2-dihydro-2-quinazolineethanamine;
2-(2-(2-azidoethyl)phenyl)-1,2-dihydro-4-quinazolinamine;
15 ethyl *N*-(4-amino-1,2-dihydroquinazolin-2-ylpropyl)carbamate;
ethyl *N*-(4-amino-1,2-dihydroquinazolin-2-ylethyl)carbamate;
ethyl *N*-(4-amino-1,2-dihydroquinazolin-2-ylmethyl)carbamate;
1-(2-thiazolylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-methoxybenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
20 1-(4-cyanobenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-nitrobenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(2-furylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-ethylbenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-chlorobenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
25 1-(2-nitrobenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(3-nitrobenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(2-methylbenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(3-methylbenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(2-thienylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;

- 1-((4-hydroxy)benzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(3-hydroxybenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-(phenylmethoxy)benzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-(4,4-dimethyloxazolin-2-yl)benzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-
5 amine;
1-(2-pyridylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-pyridylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(3-pyridazinylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(3,5-dimethylbenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
10 1-(3-fluoro-4-methylbenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(3,5-difluorobenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-(1,2,3-thiadiazol-4-yl)benzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-bromobenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-iodobenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
15 1-(4-(trifluoromethyl)benzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-(methanesulphonyl)benzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-fluorobenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(5-bromo-2-furylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-([1,1'-biphenyl]-4-ylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
20 1-(5-chloro-2-thienylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(3-pyridylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-(aminosulphonyl)benzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-methylbenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(3-amino-4-chlorobenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
25 1-((2-(trifluoromethyl)phenyl)acetyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
methyl 4-(4'-aminospiro[piperidine-4,2'(1' H)-quinazoline]-1-ylcarbonyl)benzoate;
1-(4-(1H-pyrrol-1-yl)benzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
4'-aminospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxamide;

- 1-(3-methyl-1,2,4-oxadiazol-5-yl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(2-thiazolyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-nitrophenylsulphonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-methoxyphenylsulphonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
5 1-(methanesulphonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(1-oxobutyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
5'-chloro-1-(4-cyanobenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
5'-chloro-1-(2-thienylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
5'-chloro-1-(2-furylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
10 1-(4-cyanobenzoyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
5'-fluoro-1-(2-fluorobenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-chlorobenzoyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-bromobenzoyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
5'-fluoro-1-(4-iodobenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
15 5'-fluoro-1-(4-nitrobenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-ethylbenzoyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
5'-fluoro-1-(4-propylbenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-butylbenzoyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-ethynylbenzoyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
20 5'-fluoro-1-((4-aminosulphonyl)benzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
5'-fluoro-1-((4-methanesulphonyl)benzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-
amine;
5'-fluoro-1-(4-(trifluoromethoxy)benzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-
amine;
25 methyl 4-(4'-Amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-ylcarbonyl)-
benzoate;
4-(4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-ylcarbonyl)-N-(2-hydroxy-
phenyl)benzamide;

- 4-(4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-ylcarbonyl)-N-(4-methoxyphenyl)benzamide;
- 5'-fluoro-1-(4-(2-thiazolyl)benzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 1-(3,4-dichlorobenzoyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 5 1-(4-chloro-3-iodobenzoyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 1-(4-cyano-3-methylbenzoyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 1-(4-cyano-3-fluorobenzoyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 5'-fluoro-1-(2-furylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 5'-fluoro-1-(2-thienylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 10 5'-fluoro-1-(3-thienylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 1-(4-bromo-2-thienylcarbonyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 1-(5-bromo-3-thienylcarbonyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 5'-fluoro-1-(5-chloro-2-thienylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 1-(5-bromo-2-thienylcarbonyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 15 5'-fluoro-1-(5-methyl-2-thienylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 1-(5-ethyl-2-thienylcarbonyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 5'-fluoro-1-(1H-pyrrol-2-ylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 5'-fluoro-1-(1-methyl-1H-pyrrol-2-ylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 20 5'-fluoro-1-(3-isoxazolylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 5'-fluoro-1-(5-isoxazolylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 5'-fluoro-1-(2-thiazolylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 5'-fluoro-1-(5-thiazolylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 1-(2-(3-bromo-2-thienyl)-5-thiazolylcarbonyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quin-
- 25 azoline]-4'-amine;
- 5'-fluoro-1-(4-isothiazolylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 5'-fluoro-1-(1,2,3-thiadiazol-4-ylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;

- 5'-fluoro-1-(4-pyridylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
5'-fluoro-1-(3-pyridylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(6-chloro-3-pyridylcarbonyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(6-cyano-3-pyridylcarbonyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
5 5'-fluoro-1-(2-pyrazinylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
5'-fluoro-1-(5-methyl-2-pyrazinylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-
amine;
5'-fluoro-1-(2-naphthylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
5'-fluoro-1-(2-benzo[b]thienylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
10 5'-fluoro-1-(6-quinolylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(1,3-benzodioxol-5-ylcarbonyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-
amine;
1-(5-benzofuroxanylcabonyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(1,3-dihydro-1,3-dioxo-2H-isoindol-5-ylcarbonyl)-5'-fluorospiro[piperidine-4,2'(1' H)-
15 quinazoline]-4'-amine;
O-ethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carbothioate;
5'-fluoro-1-(2-thienyl)iminomethylspiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-((4-cyanophenyl)thioxomethyl)-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-
amine;
20 5'-fluoro-1-(trifluoroacetyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
5'-fluoro-1-(4-phenoxybutanoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
3-(methanesulphonyl)propyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-
carboxylate;
5'-fluoro-1'-methyl-1-(2-thienylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-
25 amine;
5'-fluoro-1'-methyl-1-(4-cyanobenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-((4-aminosulphonyl)benzoyl)-5'-fluoro-1'-methyl-spiro[piperidine-4,2'(1' H)-quin-
azoline]-4'-amine;
1-(4-cyanobenzoyl)-5',8'-difluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;

- 1-(4-chlorobenzoyl)-5',8'-difluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
5',8'-difluoro-1-(2-thienylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
5',8'-difluoro-1-(2-pyrazinylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(6-chloro-3-pyridylcarbonyl)-5',8'-difluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-
5 amine;
1-(6-cyano-3-pyridylcarbonyl)-5',8'-difluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-
amine;
1-(4-cyanobenzoyl)-5',7'-difluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
5',7'-difluoro-1-(2-thienylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
10 1-(4-cyanobenzoyl)-5'-methoxyspiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-bromobenzoyl)-5'-hydroxyspiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(4-cyanobenzoyl)-5'-hydroxyspiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
ethyl 4'-aminospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
1-acetylspiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
15 methyl 4'-aminospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
1-methylethyl 4'-aminospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
1-benzoylspiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
ethyl 4'-amino-5'-chlorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
ethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
20 1-benzoyl-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
ethyl 4'-amino-5'-hydroxyspiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
ethyl 4'-amino-5'-methoxyspiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
ethyl 4'-amino-5',8'-difluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
ethyl 4'-amino-5',7'-difluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
25 ethyl 4'-amino-8'-chloro-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
ethyl 4'-amino-5'-fluoro-1'-methylspiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
ethyl 4'-aminospiro[piperidine-3,2'(1' H)-quinazoline]-1-carboxylate;
ethyl 4'-aminospiro[pyrrolidine-3,2'(1' H)-quinazoline]-1-carboxylate;

- propyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
methyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
2-methylpropyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
cyclopentyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
5 2-methoxyethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
S-ethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carbothioate;
2-phenoxyethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
1-methylethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
butyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
10 pentyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
hexyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
cyclobutyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
prop-2-yn-1-yl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
but-3-yn-1-yl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
15 pent-4-yn-1-yl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
hex-5-yn-1-yl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
2,2,2-trifluoroethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
4,4,4-trifluorobutyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
20 oxylate;
3-chloropropyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
4-chlorobutyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
5-chloropentyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
6-chlorohexyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
25 2-cyanoethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
2-(methylthio)ethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
3-(methylthio)propyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;

- oxylate;
- 2-phenylethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 3-phenylpropyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 4-phenylbutyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 5 2-(2-pyridyl)ethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 2-(3-pyridyl)ethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 3-(2-pyridyl)propyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 10 oxylate;
- 2-(2-pyridylthio)ethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 2-(phenylthio)ethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 15 2-(phenylamino)ethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 2-(*N*-ethyl-*N*-phenylamino)ethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 2-(4-chlorophenoxy)ethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 20 1-carboxylate;
- 2-benzofuranylmethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 3-phenoxypropyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 2-(2-thienyl)ethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 25 oxylate;
- 3-(2-thienyl)propyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 4-(2-thienyl)butyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;

- 2-(phenylmethoxy)ethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 3-(1,3-dihydro-1,3-dioxo-2H-isoindol-2-yl)propyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 5 3-(2-oxo-1(2H)-pyridyl)propyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 2-(phenylmethoxy)phenyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 5-bromo-2-methoxyphenylmethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 10 azoline]-1-carboxylate;
- 2-(4-methyl-5-thiazolyl)ethyl 4'-amino-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- phenyl 4'-aminospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 4-chlorobutyl 4'-amino-5',8'-difluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 15 oxylate;
- 4-chlorobutyl 4'-amino-5'-fluoro-1'-methylspiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxylate;
- 4'-amino-5'-fluoro-1-(1H-imidazol-1-ylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline];
- 5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 20 spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 1-(phenylmethyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 1-(phenylmethyl)spiro[piperidine-3,2'(1' H)-quinazoline]-4'-amine;
- 1-(phenylmethyl)spiro[pyrrolidine-3,2'-[1' H]-quinazoline]-4'-amine;
- 5'-fluoro-1-(phenylmethyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 25 5'-fluoro-1-(1-pyrrolidinylcarbonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
- 4'-amino-N-ethyl-5'-fluorospiro[piperidine-4,2'(1' H)-quinazoline]-1-carboxamide;
- ethyl 4'-amino-5'-fluorospiro[azetidine-3,2'(1' H)-quinazoline]-1-carboxylate;
- phenylmethyl 4'-amino-5'-fluorospiro[azetidine-3,2'(1' H)-quinazoline]-1-carboxylate;
- 5'-fluorospiro[azetidine-3,2'(1' H)-quinazoline]-4'-amine;

- 5'-fluoro-1-(2-thienylcarbonyl)spiro[azetidine-3,2'(1' H)-quinazoline]-4'-amine;
1-(3,5-dimethylisoxazol-4-yl)sulphonyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
2-ethynyl-1,2-dihydro-4-quinazolinamine;
2-(2-(2-aminoethyl)phenyl)-1,2-dihydro-4-quinazolinamine;
5 1-(4-aminobenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine;
1-(3-aminobenzoyl)spiro[piperidine-4,2'(1' H)-quinazoline]-4'-amine; or
4-(4'-aminospiro[piperidine-4,2'(1' H)-quinazoline]-1-ylcarbonyl)benzoic acid;
or a pharmaceutically acceptable salt, enantiomer or tautomer thereof.
- 10 16. A combination as claimed in any one of claims 1 to 15, comprising as a COX-2 inhibitor, at least one of Celecoxib, Meloxicam, L-745337, MK-966, L-768277, GR-253035, JTE-522, RS-57067-000, SC-58125, SC-078, PD-138387, NS-398, Flosulide and PD-164387, or pharmaceutically acceptable salts, enantiomers or tautomers thereof.
- 15 17. A combination as claimed in any one of claims 1 to 15, wherein the COX-2 inhibitor is Celecoxib or MK-966, or a pharmaceutically acceptable salt, enantiomer or tautomer thereof.
18. A combination according to any one of claims 1 to 17 for use in therapy.
- 20 19. Use of the combination as claimed in any one of claims 1 to 17 in the manufacture of a medicament for the treatment or prophylaxis of inflammatory disease.
- 25 20. A method of treatment of an inflammatory disease in a person suffering from or susceptible to such a disease, which method comprises administering to the person a therapeutically effective amount of a combination according to any one of claims 1 to 17.

21. A pharmaceutical composition comprising a combination according to any one of claims 1 to 17 in association with a pharmaceutically acceptable adjuvant, diluent or carrier.

INTERNATIONAL SEARCH REPORT

International application No.

PCT/SE 99/01144

A. CLASSIFICATION OF SUBJECT MATTER

IPC6: A61K 31/505, A61K 31/415

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC6: A61K

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

SE,DK,FI,NO classes as above

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	WO 9714686 A1 (ASTRA PHARMACEUTICALS LIMITED), 24 April 1997 (24.04.97) --	1-21
P,A	STN International, File MEDLINE, Medline accession no. 1999000472, Hamilton L C et al: Interactions between inducible isoforms of nitric oxide synthase and cyclo-oxygenase in vivo: investigations using the selective inhibitors, 1400W and celecoxib"; & British Journal of Pharmacology, (1998 Sep) 125 (2) 335-40 -- -----	1-21

☐ Further documents are listed in the continuation of Box C.☒ See patent family annex.

* Special categories of cited documents:

"A" document defining the general state of the art which is not considered to be of particular relevance

"E" earlier document but published on or after the international filing date

"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)

"O" document referring to an oral disclosure, use, exhibition or other means

"P" document published prior to the international filing date but later than the priority date claimed

"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention

"X" document of particular relevance: the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone

"Y" document of particular relevance: the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art

"&" document member of the same patent family

Date of the actual completion of the international search

4 October 1999

Date of mailing of the international search report

22 -10- 1999

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INTERNATIONAL SEARCH REPORT

International application No.
PCT/SE 99/01144

Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)

This international search report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. ☒ Claims Nos.: 20
because they relate to subject matter not required to be searched by this Authority, namely:
see next sheet
2. ☐ Claims Nos.:
because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:

Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

1. ☐ As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.
2. ☐ As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. ☐ As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:
4. ☐ No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

Remark on Protest

- ☐ The additional search fees were accompanied by the applicant's protest.
☐ No protest accompanied the payment of additional search fees.